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Substitute for Form 1449/A/PTO

# INFORMATION DISCLOSURE STATEMENT BY APPLICANT

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Application No.	10/077,142
Filing Date	02/15/2002
First Named Inventor	D'Amato et al.
Group Art Unit	1616
Examiner Name	Badio, Barbara P.
Attorney Docket Number	05213-3000 (43170-269288)

Sheet 1 of 15

## U.S. PATENT DOCUMENTS

Examiner Initials	Cite No. <sup>1</sup>	U.S. Patent Document		Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
		Number <sup>2</sup>	Kind Code <sup>3</sup> (if known)			
	1	3,166,577		Ringold et al.	01/19/1965	
	2	3,562,260		De Ruggieri et al.	02/09/1971	
	3	4,552,758		Murphy et al.	11/12/1985	
	4	4,808,402		Leibovich et al.	02/28/1989	

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**FOREIGN PATENT DOCUMENTS**

Examiner Initials	Cite No. <sup>1</sup>	Foreign Patent Document		Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear	T <sup>2</sup>
		Office <sup>3</sup>	Number <sup>4</sup>	Kind Code <sup>5</sup> (if known)			
	5	WO	88/08002	A1	The Upjohn Company	10/20/1988	
	6	WO	93/19746	A1	ORION-YH-TYMA OY	10/14/1993	

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Sheet	3	of	15	Attorney Docket Number	05213-3000 (43170-269288)
<b>OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS</b>					
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	7	Author: Title: Publi: Research Plus Catalog Vol/Iss: Pages: 50-58 Date: 1993			
	8	Author: Title: Registry Number: 56933-77-8 Publi: Chemical Abstracts Vol/Iss: Pages: Date:			
	9	Author: Title: Registry Number: 56933-78-9 Publi: Chemical Abstracts Vol/Iss: Pages: Date:			
	10	Author: Title: Registry Number: 57380-15-1 Publi: Chemical Abstracts Vol/Iss: Pages: Date:			
	11	Author: Title: Registry Number: 71782-94-0 Publi: Chemical Abstracts Vol/Iss: Pages: Date:			
	12	Author: Title: Registry Number: 71782-95-1 Publi: Chemical Abstracts Vol/Iss: Pages: Date:			
	13	Author: Title: Registry Number: 101277-11-6 Publi: Chemical Abstracts Vol/Iss: Pages: Date:			

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	14	Author: Title: Registry Number: 101429-40-7 Publi: Chemical Abstracts Vol/Iss: Pages: Date:			
	15	Author: Title: Registry Number: 162853-20-5 Publi: Chemical Abstracts Vol/Iss: Pages: Date:			
	16	Author: ABOULWafa et al. Title: Synthesis and evaluation for uterotrophic and antiimplantation activities of 2-substituted estradiol derivatives Publi: Steroids Vol/Iss: 57 Pages: 199-204 Date: 04/1992			
	17	Author: ALGIRE, G.H. et al. Title: Vascular reactions of normal and malignant tumors in vivo. I. Vascular reactions of mice to wounds and to normal and neoplastic transplants Publi: Journal of the National Cancer Institute Vol/Iss: 6 Pages: 73-85 Date: 08/1945			
	18	Author: ALIEV et al. Title: 54929q Synthesis of cycloalkyl derivatives of dihydric phenols and their ethers Publi: Chemical Abstracts Vol/Iss: 72 Pages: 370 Date: 1970			
	19	Author: BARNES et al. Title: Tumor Necrosis Factor Production in Patients with Leprosy Publi: Infection and Immunity Vol/Iss: 60 (4) Pages: 1441-1446 Date: 04/1992			
	20	Author: BINDRA et al. Title: Studies in Antifertility Agents. 8. Seco Steroids. 2. 5,6-Secoestradiol and Some Related Compounds Publi: Journal of Medicinal Chemistry Vol/Iss: 18 (9) Pages: 921-925 Date: 1975			

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	21	Author: BLICKENSTAFF et al. Title: SYNTHESIS OF SOME ANALOGS OF ESTRADIOL Publ: Steroids Vol/Iss: 46 (4,5) Pages: 889-902 Date: 10/1985	
	22	Author: BOYCE et al. Title: Some Preliminary Synthetical Studies with 5,6,7,8-Tetra-hydro-8-methylindane -1,5-dione Publ: Unknown Vol/Iss: Pages: 4547-4553 Date: 1960	
	23	Author: BRANDI et al. Title: Bone endothelial cells as estrogen targets (Abstract only) Publ: Calcif. Tissue Int. Vol/Iss: 53 (5) Pages: 312-317 Date: 1993	
	24	Author: BREM, H. et al. Title: Interstitial chemotherapy with drug polymer implants for the treatment of recurrent gliomas Publ: Journal of Neurosurgery Vol/Iss: 74 Pages: 441-446 Date: 03/01/91	
	25	Author: CAMBIE et al. Title: Aromatic Steroids. Part I. Oxidation Products of 3-Methoxyestra-1,3,5(10)-triene-17 $\beta$ -yl Acetate Publ: J. Chem. Soc. Vol/Iss: Pages: 2603-2608 Date: 1968	
	26	Author: COREY et al. Title: Applications of N,N-Dimethylhydrazones to Synthesis. Use in Efficient, Positionally and Stereochemically Selective C-C Bond Formation; Oxidative Hydrolysis to Carbonyl Compounds Publ: Tetrahedron Letters Vol/Iss: 1 Pages: 3-6 Date: 1976	
	27	Author: COREY et al. Title: Facile Conversion of N,N-Dimethylhydrazones to Carbonyl Compounds by Cupric Ion-Catalyzed Hydrolysis Publ: Tetrahedron Letters Vol/Iss: 41 Pages: 36678-3668 Date: 1976	

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	28	Author: CRABBE, P. Title: Cotton effect of the styrene chromophore (Abstract only) Publ: Chem. Ind. Vol/Iss: 27 Pages: 917-918 Date: 1969			
	29	Author: DURANI et al. Title: Seco-Oestradiols and Some Non-Steroidal Oestrogens: Structural Correlates of Oestrogenic Action Publ: Journal of Steroid Biochemistry Vol/Iss: 11 Pages: 67-77 Date: 1979			
	30	Author: EDER et al. Title: Synthese von Ostradiol (in German - No translation available) Publ: Chem. Ber. Vol/Iss: 109 Pages: 2948-2953 Date: 1976			
	31	Author: EMONS et al. Title: Modulation der hypophysären Sekretion von Luteinisierendem Hormon (LH) durch Östrogene Publ: FOCUS MHL Vol/Iss: 3 Pages: 221-228 Date: 1986			
	32	Author: FANCHENKO et al. Title: Characteristics of the guinea pig uterus estrogen receptor system (Abstract only) Publ: Byull. Eksp. Biol. Med. Vol/Iss: 85 (4) Pages: 467-470 Date: 1978			
	33	Author: FEVIG et al. Title: A Short, Stereoselective Route to 16 $\alpha$ -(Substituted-alkyl)estradiol Derivatives Publ: Journal of Organic Chemistry Vol/Iss: 52 Pages: 247-251 Date: 1987			
	34	Author: FIELD et al. Title: Effect of Thalidomide on the Graft versus Host Reaction Publ: Nature Vol/Iss: 211 (5055) Pages: 1308-1310 Date: 09/17/66			

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	91	Author: VICENTE et al. Title: In Vitro Activity of Thalidomide Against Mycobacterium avium Complex Publ: Archives of Internal Medicine Vol/Iss: 153 (4) Pages: 534 Date: 02/22/93	
	92	Author: WANG et al. Title: Photoaffinity labeling of human placental estradiol 17.β-dehydrogenase with 2- and 4-azidoestrone, 2- and 4-azidoestradiol (Abstract only) Publ: Shengwu Huaxue Zazhi Vol/Iss: 8 (6) Pages: 715-718 Date: 1992	
	93	Author: WEIDNER, N. et al. Title: Tumor angiogenesis: A New Significant and Independent Prognostic Indicator in Early-Stage Breast Carcinoma Publ: Journal of the National Cancer Institute Vol/Iss: 84 Pages: 1875-1887 Date: 12/16/92	
	94	Author: WEIDNER, N. et al. Title: Tumor Angiogenesis Correlates with Metastasis in Invasive Prostate Carcinoma Publ: American Journal of Pathology Vol/Iss: 143 (2) Pages: 401-409 Date: 08/1993	
	95	Author: WEIDNER, N. et al. Title: Tumor Angiogenesis and Metastasis-Correlation in Invasive Breast Carcinoma Publ: New England Journal of Medicine Vol/Iss: 324 (1) Pages: 1-8 Date: 01/03/91	
	96	Author: WELSCH et al. Title: Staphylostatic Activity of Some New Diphenols, Napthols, and Chalcones (Abstract only) Publ: Experientia Vol/Iss: 11 Pages: 350-351 Date: 1955	
	97	Author: WHITE et al. Title: Treatment of Pulmonary Hemangiomas with Recombinant Interferon Alfa-2a Publ: The New England Journal of Medicine Vol/Iss: 32 (18) Pages: 1197-1200 Date: 05/04/89	

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	77	Author: NEWKOME et al. Title: Synthesis of Simple Hydrazones of Carbonyl Compounds by an Exchange Reaction Publ: Journal of Organic Chemistry Vol/Iss: 31 Pages: 677-681 Date: 03/1966	
	78	Author: NGUYEN, M. et al. Title: Elevated Levels of the Angiogenic Peptide Basic Fibroblast Growth Factor in Urine of Bladder Cancer Patients Publ: Journal of the National Cancer Institute Vol/Iss: 85 (3) Pages: 241-242 Date: 02/03/93	
	79	Author: NUMAZAWA et al. Title: Novel and Regiospecific Synthesis of 2-Amino Estrogens via Zincke Nitration Publ: Steroids Vol/Iss: 41 (5) Pages: 675-682 Date: 1983	
	80	Author: PERT et al. Title: Preparations of 2,4-disubstituted estradiols (Abstract only) Publ: Australian Journal of Chemistry Vol/Iss: 42 (3) Pages: 421-432 Date: 1989	
	81	Author: PETERS et al. Title: 17-Desoxy Estrogen Analogues Publ: Journal of Medicinal Chemistry Vol/Iss: 32 (7) Pages: 1642-1652 Date: 1989	
	82	Author: PFEIFFER et al. Title: Are catechol estrogens obligatory mediators of estrogen action in the central nervous system? I. Characterization of pharmacological probes with different receptor binding affinities and catechol estrogen formation rates (Abstract only) Publ: Journal of Endocrinology Vol/Iss: 110 (3) Pages: 489-497 Date: 1986	
	83	Author: POWELL et al. Title: Investigation and Treatment of Orogenital Ulceration; studies on a Possible Mode of Action of Thalidomide Publ: British Journal of Dermatology Vol/Iss: 113 Supp. 28 Pages: 141-144 Date: 07/1985	

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	84	Author: SHISHKINA et al. Title: Synthesis and properties of condensed heterocyclic derivatives of estra-4, 9-dien-17.β-ol-3-one (Abstract only) Publ: Khim.-Farm. Zh. Vol/Iss: 8 (1) Pages: 7-11 Date: 1974			
	85	Author: SIDKY et al. Title: Inhibition of Angiogenesis by Interferons: Effects on Tumor- and Lymphocyte-induced Vascular Responses Publ: Cancer Research Vol/Iss: 47 Pages: 5155-5161 Date: 10/01/87			
	86	Author: SIRACUSA et al. Title: The effect of microtubule- and microfilament-disrupting drugs on preimplantation mouse embryos (Abstract only) Publ: Journal of Embryology and Experimental Morphology Vol/Iss: 60 Pages: 71-82 Date: 12/1980			
	87	Author: SPYRIOUNIS et al. Title: Copper (II) complex of an estradiol derivative with potent antiinflammatory properties (Abstract only) Publ: Arch. Pharm. Vol/Iss: 324 (9) Pages: 533-536 Date: 1991			
	88	Author: SRIVASTAVA, A. et al. Title: The Prognostic Significance of Tumor Vascularity in Intermediate-Thickness (0.76-4.0 mm Thick) Skin Melanoma Publ: American Journal of Pathology Vol/Iss: 133 (2) Pages: 419-424 Date: 11/1988			
	89	Author: STARKOV et al. Title: Mono- and Dialkylation of Guaiacol by Olefins on KU-2 Cation Exchanger (Abstract only) Publ: Zhurnal Prikladnoi Khimii Vol/Iss: 41 (3) Pages: 688-690 Date: 1968			
	90	Author: TAYLOR, S. et al. Title: Protamine is an Inhibitor of Angiogenesis Publ: Nature Vol/Iss: 297 Pages: 307-312 Date: 05/27/82			

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	70	Author: LOOZEN et al. Title: An approach to the synthesis of 7.beta.-amino estrogens (Abstract only) Publ: Recl. J.R. Neth.Chem. Soc. Vol/Iss: 102 (10) Pages: 433-437 Date: 1983	
	71	Author: MARO et al. Title: Mechanism of Polar Body Formation in the Mouse Oocyte: An Interaction Between the Chromosomes, the Cytoskeleton and the Plasma Membrane Publ: Journal of Embryology and Experimental Morphology Vol/Iss: 92 Pages: 11-32 Date: 1986	
	72	Author: MICHEL et al. Title: Inhibition of synaptosomal high-affinity uptake of dopamine and serotonin by estrogen agonists and antagonists (Abstract only) Publ: Biochem. Pharmacol. Vol/Iss: 36 (19) Pages: 3175-3180 Date: 1987	
	73	Author: MORISAKI et al. Title: Steroids. LI. Aromatization reaction of the cross-conjugated dienone system by Zinc 9. (Abstract only) Publ: Chem. Pharm. Bull. Vol/Iss: 14 (8) Pages: 866-872 Date: 1966	
	74	Author: NAAFS et al. Title: Thalidomide Therapy An Open Trial Publ: International Journal of Dermatology Vol/Iss: 24 (2) Pages: 131-134 Date: 03/1985	
	75	Author: NAMBARA et al. Title: Microbial transformation products derived from steroids. I. Synthesis of 1,2- and 3-dimethoxy-4-methylestratrienes (Abstract only) Publ: Chem. Pharm. Bull. Vol/Iss: 20 (2) Pages: 336-342 Date: 1972	
	76	Author: NAMBARA et al. Title: Synthesis of 16β-Oxygenated Catechol Estrogen Methyl Ethers, New and Potential Metabolites Publ: Chemical & Pharmaceutical Bulletin Vol/Iss: 23 (7) Pages: 1613-1616 Date: 07/1975	

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	63	Author: KNIGHTON, D. et al. Title: Avascular and Vascular Phases of Tumour Growth in the Chick Embryo Publ: British Journal of Cancer Vol/Iss: 35 Pages: 347-356 Date: 1977				
	64	Author: KOLÉ et al. Title: Studies in Antifertility Agents. 11. Secosteroids. 5. Synthesis of 9,11-Secoestradiol Publ: Journal of Medicinal Chemistry Vol/Iss: 18 (7) Pages: 765-766 Date: 1975				
	65	Author: KOVACS et al. Title: Steroids. XXIII. Synthesis of 2- and 4-hydroxy and 2,4-dihydroxy derivatives of estrone and estradiol (Abstract only) Publ: Acta Phys. Chem. Vol/Iss: 19 (3) Pages: 287-290 Date: 1973				
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<sup>1</sup>Unique citation designation number. <sup>2</sup>See attached Kinds of U.S. Patent Documents. <sup>3</sup>Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). <sup>4</sup>For Japanese patent document, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. <sup>5</sup>Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. <sup>6</sup>Applicant is to place a check mark here if English language translation is attached.

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<b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b> (use as many sheets as necessary)				Application No.	10/077,142
				Filing Date	02/15/2002
				First Named Inventor	D'Amato et al.
				Group Art Unit	1616
				Examiner Name	Badio, Barbara P.
Sheet	7	of	15	Attorney Docket Number	05213-3000 (43170-269288)
<b>OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS</b>					
Examiner Initials	Cite No.	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published			T*
	35	Author: FIESER et al. Title: N-Methylformanilide Publ: Organic Synthesis Collective Volume 3 Vol/Iss: 3 Pages: 590-591 Date: 1955			
	36	Author: FLOHE et al. Title: Studies on the Hypothetical Relationship of Thalidomide-induced Embryopathy and Collagen Biosynthesis Publ: Arzneimittel/Forschung (Germany West) Vol/Iss: 31 (2) Pages: 315-320 Date: 01/01/81			
	37	Author: FOLKMAN et al. Title: Angiogenesis Inhibition and Tumor Regression Caused by Heparin or a Heparin Fragment in the Presence of Cortisone Publ: Science Vol/Iss: 221 Pages: 719-725 Date: 08/19/83			
	38	Author: FOLKMAN, J. Title: Tumor Angiogenesis: Therapeutic Implications Publ: New England Journal of Medicine Vol/Iss: 285 (21) Pages: 1182-1186 Date: 11/18/71			
	39	Author: FOLKMAN, J. et al. Title: Induction of Angiogenesis During the Transition from Hyperplasia to Neoplasia Publ: Nature Vol/Iss: 339 Pages: 58-61 Date: 05/04/89			
	40	Author: FOLKMAN, J. et al. Title: Tumor Behavior in Isolated Perfused Organs in vitro Growth and Metastases of Biopsy Material in Rabbit Thyroid and Canine Intestinal Segment Publ: Annals of Surgery Vol/Iss: 164(3) Pages: 491-502 Date: 09/01/66			
	41	Author: GANDHI et al. Title: Mannich reaction of estrone Publ: Journal of Indian Chem. Soc. Vol/Iss: 39 Pages: 306-308 Date: 1962			

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	42	Author: GASLINI et al. Title: Reaction of Eugenol with Synthesis Gas. Synthesis of 5,6,7,8-Tetrahydro-3-methoxy-2-naphthol Publ: Journal of Organic Chemistry Vol/Iss: 29 (5) Pages: 1177-1180 Date: 05/1964			
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	44	Author: GIMBRONE, M.A. et al. Title: Tumor dormancy <i>in vivo</i> by Prevention of Neovascularization Publ: Journal of Experimental Medicine Vol/Iss: 136 Pages: 261-276 Date: 1972			
	45	Author: GONZALEZ et al. Title: Synthesis and Pharmacological Evaluation of 8αEstradiol Derivatives Publ: Steroids Vol/Iss: 40 (2) Pages: 171-187 Date: 9/1982			
	46	Author: GROSS, J.L. et al. Title: Modulation of Solid Tumor Growth <i>in vivo</i> by bFGF (Abstract only) Publ: Proceedings of the American Association of Cancer Research Vol/Iss: 31 Pages: 79 Date: 03/90			
	47	Author: GUNZLER, V. Title: Thalidomide-A Therapy for the Immunological Consequences of HIV Infection? Publ: Medical Hypothesis Vol/Iss: 30 (2) Pages: 105-109 Date: 10/89			
	48	Author: GUPTA et al. Title: Antifertility Agents. XIV. Secosteroids. VII. Synthesis of 2α and 2β, 6β-dimethyl-3β-(p-hydroxyphenyl)-trans-bicyclo[4.3.0]nonan-7-ones and some related compounds (Abstract only) Publ: Indian Journal of Chemistry Vol/Iss: 13 (7) Pages: 759-760 Date: 1975			

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	49	Author: GUPTA et al. Title: Studies in Antifertility Agents. Part XVIII. 2α,6β-Diethyl-3β-(p-hydroxyphenyl)-trans-bicyclo[4.3.0]nonan-7β-ol and 6β-methyl-3β-(p-hydroxyphenyl)-2α-propyl-trans-bicyclo[4.3.0]nonan-7β-ol (Abstract only) Publi: Indian Journal of Chemistry Vol/Iss: 19B (10) Pages: 886-890 Date: 1980			
	50	Author: GUTIERREZ-RODRIGUEZ et al. Title: Treatment of Refractory Rheumatoid Arthritis - The Thalidomide Experience Publi: The Journal of Rheumatology Vol/Iss: 16 (2) Pages: 158-163 Date: 02/89			
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	52	Author: HAHNEL et al. Title: THE SPECIFICITY OF THE ESTROGEN RECEPTOR OF HUMAN UTERUS Publi: Journal of Steroid Biochemistry Vol/Iss: 4 Pages: 21-31 Date: 1973			
	53	Author: HANDLEY et al. Title: Chronic bullous disease of childhood and ulcerative colitis Publi: British Journal of Dermatology Vol/Iss: 127 (40) Pages: 67-68 Date: 07/01/92			
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	55	Author: HOLKER et al. Title: The Reactions of Estrogens with Benzeneseleninic Anhydride and Hexamethyldisilazane Publi: J. Chem. Soc. Perkin Trans. Vol/Iss: 1 Pages: 1915-1918 Date: 1982			

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	56	Author: HORI, A. et al. Title: Suppression of Solid Tumor Growth by Immunoneutralizing Monoclonal Antibody Against Human Basic Fibroblasts Growth Factor Publ: Cancer Research Vol/Iss: 51 Pages: 6180-6184 Date: 11/15/91			
	57	Author: IMAMURA et al. Title: Method for Manufacture of Dihydric Phenols (Abstract only) Publ: USPATEFULL 76:20259 US 3,950,437 Vol/Iss: Pages: Date: 04/13/1976			
	58	Author: INGBER, D. et al. Title: Synthetic analogues of fumagillin that inhibit angiogenesis and suppress tumor growth Publ: Nature Vol/Iss: 348 Pages: 555-557 Date: 12/06/90			
	59	Author: IRIARTE et al. Title: Steroids (XCIV). Synthesis of 2-methyl and 1,2-dimethyl estrogens (Abstract only) Publ: Tetrahedron Vol/Iss: 3 Pages: 28-36 Date: 1958			
	60	Author: JHINGRAN et al. Title: Studies in Antifertility Agents - Part XLI: Secosteroids-x: Syntheses of Various Stereoisomers of (+)-2,6β-diethyl-7α-ethynyl-3-(p-hydroxyphenyl)-trans-bicyclo [4.3.0]nonan-7β-ol. Publ: Steroids Vol/Iss: 42 (6) Pages: 627-634 Date: 1983			
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	62	Author: KIM, K.J. et al. Title: Inhibition of Vascular Endothelial Growth Factor-induced Angiogenesis Suppresses Tumor Growth In Vivo Publ: Nature Vol/Iss: 362 Pages: 841-844 Date: 04/29/93			

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